

**ABSTRACT**

The present invention is directed towards novel cyclic phosphoramidate prodrugs of alcohol-, amine-, and thiol-containing drugs, their preparation, their synthetic intermediates, and their uses. Another aspect of the invention is the use of the prodrugs to treat diseases that benefit from enhanced drug distribution to the liver and like tissues and cells that express cytochrome P450, including hepatitis, cancer, liver fibrosis, malaria, other viral and parasitic infections, and metabolic diseases where the liver is responsible for the overproduction of the biochemical end product, e.g. glucose (diabetes); cholesterol, fatty acids and triglycerides (hyperlipidemia) (atherosclerosis) (obesity). In one aspect, the invention is directed towards the use of the prodrugs to enhance oral drug delivery. In another aspect, the prodrugs are used to prolong pharmacodynamic half-life of the drug. In addition, the prodrug methodology of the current invention is used to achieve sustained delivery of the parent drug. In another aspect, the prodrugs are used to increase the therapeutic index of the drug. In another aspect of the invention, a method of making these prodrugs is described. In another aspect, the prodrugs are also useful in the delivery of diagnostic imaging agents to the liver.